

IN THE CLAIMS

Please amend the pending claims as follows:

1-61. (Previously canceled)

62. (Original) A method for screening for primary N-hydroxylamines which reduce oxidative damage to, or delay senescence of cells, comprising the steps of:

contacting cells with a candidate primary N-hydroxylamine under conditions whereby, but for the presence of the hydroxylamine, the cells present a reference amount of oxidative damage or senescence;

detecting post-treatment amounts of oxidative damage or senescence of the cells;

wherein a lesser amount of post-treatment than reference amounts of oxidative damage or senescence indicates that the hydroxylamine reduces oxidative damage or delays senescence of the cells.

63. (Previously presented) A method according to claim 62, wherein the cells are in an animal and the hydroxylamine is administered to the animal in a dosage from 100ug to 1g.

64. (Previously presented) A method according to claim 62, wherein the hydroxylamine has the general formula,

$\text{NHOHCR}_1\text{R}_2\text{R}_3$,

wherein R_1 , R_2 and R_3 are independently selected from: hydrogen, substituted or unsubstituted (C1-C10) alkyl, alkenyl, alkynyl, aryl, oxyl, acyl, carboxyl, amino, nitro, nitroso, oxime, hydrazone, azo, thiol, sulfonyl and halide.

65. (Currently amended) A method according to claim 64, wherein at least one R of R_1 , R_2 and R_3 is selected from unsubstituted (C1-C10) alkyl, alkenyl and alkynyl.

66. (Currently amended) A method according to claim 64, wherein at least one R of R_1 , R_2 and R_3 is selected from unsubstituted (C1-C18) alkyl, cycloalkyl, alkenyl and alkynyl, and the R is selected from: $\text{CH}_3\text{-(CH}_2\text{)}_{n1}$, $(\text{CH}_3\text{-(CH}_2\text{)}_{n2}\text{-})_2\text{CH}$, $(\text{CH}_3\text{-(CH}_2\text{)}_{n2}\text{-})_3$, cyclopentyl, cyclohexyl, $(\text{CH}_2$

$=\text{CH}-\text{CH}_2)_{n3}$ and $(\text{CH} \equiv \text{C}-\text{CH}_2)_{n3}$, wherein $n1 = 1$ to 18 , $n2 = 1$ to 17 and $n3 = 1$ to 3 .

67. (Currently amended) A method according to claim 64, wherein at least one R of R_1 , R_2 and R_3 is selected from hydrogen, unsubstituted (C1-C10) alkyl, alkenyl and alkynyl, and the hydroxylamine is selected from:

N-methylhydroxylamine,	N-(n-decahexyl)hydroxylamine,
N-ethylhydroxylamine,	N-(n-decaoctyl)hydroxylamine,
N-n-propylhydroxylamine,	N-isopropylhydroxylamine,
N-(n-butyl) hydroxylamine,	N-sec-butylhydroxylamine,
N-(n-pentyl)hydroxylamine,	N-tert-butylhydroxylamine,
N-(n-hexyl)hydroxylamine,	N-cyclohexylhydroxylamine,
N-(n-heptyl)hydroxylamine,	N-cyclopentylhydroxylamine,
N-(n-octyl)hydroxylamine,	N-(2-propene)hydroxylamine,
N-(n-nonyl)hydroxylamine,	N-(3-butene)hydroxylamine,
N-(n-decyl)hydroxylamine,	N-(2-propyne)hydroxylamine and
N-(n-dodecyl)hydroxylamine,	N-(3-butyne)hydroxylamine.

68. (Currently amended) A method according to claim 64, wherein at least one R of R_1 , R_2 and R_3 is substituted or unsubstituted aryl.

69. (Currently amended) A method according to claim 64, wherein at least one R of R_1 , R_2 and R_3 is substituted or unsubstituted aryl, and the R is selected from: mono, di, or tri methyl, methoxy, halo, nitro, amino, hydroxyl and substituted or unsubstituted phenyl, naphthyl, anthryl, phenanthryl, pyridyl, quinolinyl, imidazolyl, benzoxazolyl, pyrrolyl, furanyl, piperidinolyl and tetrahydrofuranyl.

70. (Currently amended) A method according to claim 64, wherein at least one R of R_1 , R_2 and R_3 is substituted or unsubstituted aryl, and the hydroxylamine is selected from:

N-benzylhydroxylamine,
N-(n-nitrobenzyl)hydroxylamine,
N-(n-methylbenzyl)hydroxylamine,
N-(n-chlorobenzyl)hydroxylamine,
N-(n-aminobenzyl)hydroxylamine,
N-(n-hydroxybenzyl)hydroxylamine,

wherein n is selected from 1, 2, 3, 4, 5 and 6.

N-(1,3-diaminobenzyl)hydroxylamine,
N-(1,3-hydroxybenzyl)hydroxylamine,
N-(2,4-diaminobenzyl)hydroxylamine,
N-(2,4-dihydroxybenzyl)hydroxylamine,
Imidazole-2-methylhydroxylamine and
Benzoxazole-2-methylhydroxylamine,

71. (Currently amended) A method according to claim 64, wherein at least one R of R₁, R₂ and R₃ is substituted or unsubstituted (C1-C18) oxyl.

72. (Currently amended) A method according to claim 64, wherein at least one R of R₁, R₂ and R₃ is substituted or unsubstituted (C1-C18) oxyl and the R is selected from: hydroxyl, hydroxyalkyl (HO-(CH₂)_{n1}), hydroxyaryl selected from benzylalcohol, phenol and naphthol, alkoxy (O-(CH₂)_{n1}) and aryloxy selected from phenoxy, benzyloxy and naphthyoxy, wherein n1=1 to 18.

73. (Currently amended) A method according to claim 64, wherein at least one R of R₁, R₂ and R₃ is substituted or unsubstituted (C1-C18)alkyl hydroxyl or arylhydroxyl and the hydroxylamine is selected from:

N-(hydroxymethyl)hydroxylamine,
N-(2-hydroxyethyl)hydroxylamine,
N-(3-hydroxypropyl)hydroxylamine,
N-(4-hydroxybutyl)hydroxylamine,
N-(6-hydroxyhexyl)hydroxylamine,
N-(12-hydroxydodecyl)hydroxylamine,

N-(methoxymethyl)hydroxylamine,
N-(methoxyethyl)hydroxylamine,
N-(methoxyisopropyl)hydroxylamine,
N-(benzyloxymethyl)hydroxylamine and
N-(4-hydroxymethylbenzyl)hydroxylamine.

74. (Currently amended) A method according to claim 64, wherein at least one R of R₁, R₂ and R₃ is substituted or unsubstituted (C1-C18) alkylcarboxyl or arylcarboxyl.

75. (Currently amended) A method according to claim 64, wherein at least one R of R₁, R₂ and R₃ is substituted or unsubstituted (C1-C18) alkyl or aryl carboxyl and the R is selected from carboxyalkyls and benzyl.

76. (Currently amended) A method according to claim 64, wherein at least one R of R₁, R₂ and R₃ is substituted or unsubstituted alkyl (C1-C18) or arylcarboxyl and the hydroxylamine is selected from:

N-(carboxymethyl)hydroxylamine,
N-(2-carboxyethyl)hydroxylamine,
N-(3-carboxypropyl)hydroxylamine,
N-(4-carboxybutyl)hydroxylamine,

N-(5-carboxypentyl) hydroxylamine,
N-(6-carboxyhexyl)hydroxylamine,
N-(4-carboxybenzyl)hydroxylamine and
N-(12-carboxydodecyl)hydroxylamine.

77. (Currently amended) A method according to claim 64, wherein at least one R of R₁, R₂ and R₃ is substituted or unsubstituted (C1-C18) ester.

78. (Currently amended) A method according to claim 64, wherein at least one R of R₁, R₂ and R₃ is substituted or unsubstituted (C1-C18) ester and the R is selected from alkyl (C1 – C18) and aryl esters.

79. (Currently amended) A method according to claim 64, wherein at least one R of R₁, R₂ and R₃ is substituted or unsubstituted alkyl (C1-C18) or arylesters and the hydroxylamine is selected from:

N-(acetyloxymethyl)hydroxylamine,
N-(acetyloxyethyl)hydroxylamine,
N-(acetyloxypropyl)hydroxylamine,
N-(propylcarbonyloxy)methylhydroxylamine,
N-(butylcarbonyloxy)methylhydroxylamine,
N-(tert-butyloxycarboxyl)methylhydroxylamine,
N-(benzyloxycarbonyl)methylhydroxylamine,
N-(phenyloxycarbonyl)methylhydroxylamine,

N-(3-pyridyloxycarbonyl)methylhydroxylamine and
N-(benzoxazol-5-carbonyloxy)methylhydroxylamine.

80. (Currently amended) A method according to claim 64, wherein at least one R of R₁, R₂ and R₃ is substituted or unsubstituted (C1-C18) carbonyl.

81. (Currently amended) A method according to claim 64, wherein at least one R of R₁, R₂ and R₃ is substituted or unsubstituted carbonyl and the R is selected from alkyl (C1 – C18) carbonyls and aryl carbonyls.

82. (Currently amended) A method according to claim 64, wherein at least one R of R₁, R₂ and R₃ is substituted or unsubstituted alkyl (C1-C18) or arylcarbonyls and the hydroxylamine is selected from:

N-(acetyl)methylhydroxylamine,	N-(phenylcarbonyl)methylhydroxylamine and
N-(ethylcarbonyl)methylhydroxylamine,	N-(benzylcarbonyl)methylhydroxylamine.
N-(butylcarbonyl)methylhydroxylamine,	

83. (Currently amended) A method according to claim 64, wherein at least one R of R₁, R₂ and R₃ is substituted or unsubstituted alkyl(C1-C18) or aryl amino.

84. (Currently amended) A method according to claim 64, wherein at least one R of R₁, R₂ and R₃ is substituted or unsubstituted alkyl (C1-C18) or aryl amino and the R is selected from primary alkyl amine selected from methylamine, ethylamine, propylamine, butylamine and hexylamine, secondary amine selected from dimethylamine, diethylamine and dipropylamine, tertiary amine selected from trimethyl and triethylamine, and quarternary amine selected from tetramethyl and tetra-ethylammonium salts.

85. (Currently amended) A method according to claim 64, wherein at least one R of R₁, R₂ and R₃ is substituted or unsubstituted alkyl(C1-C18) or aryl amine and the hydroxylamine is selected from:

N-aminomethylhydroxylamine,
N-(2-aminoethyl)hydroxylamine,
N-(N-methylamino)methylhydroxylamine,
N-(N,N-dimethylamino)methylhydroxylamine,
N-(N,N,N-trimethylammonium)methylhydroxylamine,
N-(3-aminopropyl)hydroxylamine,
N-(6-aminohexyl)hydroxylamine,
N-(4-aminobenzyl)hydroxylamine,
Hydroxylamine-1-methylpyridinium and
Hydroxylamine-1-methylquinolinium.

86. (Currently amended) A method according to claim 64, wherein at least one R of R₁, R₂ and R₃ is substituted or unsubstituted (C1-C18) alkyl or aryl nitro.

87. (Currently amended) A method according to claim 64, wherein at least one R of R₁, R₂ and R₃ is substituted or unsubstituted alkyl(C1-C18) or aryl nitro and the R is selected from alkynitro selected from nitromethyl, nitroethyl, nitropropyl, nitrobutyl, nitropentyl, nitrohexyl and nitrobenzyl, and arynitro selected from nitrophenyl and nitronaphthyl.

88. (Currently amended) A method according to claim 64, wherein at least one R of R₁, R₂ and R₃ is substituted or unsubstituted alkyl (C1-C18) or aryl nitro and the hydroxylamine is selected from:

N-(nitromethyl)hydroxylamine,	N-(5-nitropentyl)hydroxylamine,
N-(2-nitroethyl)hydroxylamine,	N-(6-nitrohexyl)hydroxylamine,
N-(3-nitropropyl)hydroxylamine,	N-(4-nitrobenzyl)hydroxylamine and
N-(4-nitrobutyl)hydroxylamine,	N-(2,4-dinitrobenzyl)hydroxylamine.

89. (Currently amended) A method according to claim 64, wherein at least one R of R₁, R₂ and R₃ is substituted or unsubstituted (C1-C18) nitroso.

90. (Currently amended) A method according to claim 64, wherein at least one R of R₁, R₂ and R₃ is substituted or unsubstituted (C1-C18) nitroso and the R is selected from aliphatic nitrosoamines and aromatic nitroso.

91. (Currently amended) A method according to claim 64, wherein at least one R of R₁, R₂ and R₃ is substituted or unsubstituted nitroso (C1-C18) and the hydroxylamine is selected from:
N-(N-methyl-N-nitroso-amino)methyl hydroxylamine,
N-(N-methyl-N-nitroso-2-amino)ethylhydroxylamine,
N-(N-methyl-N-nitroso-3-amino)propylhydroxylamine and
N-(p-nitroso)benzylhydroxylamine.

92. (Currently amended) A method according to claim 64, wherein at least one R of R₁, R₂ and R₃ is substituted or unsubstituted oxime.

93. (Currently amended) A method according to claim 64, wherein at least one R of R₁, R₂ and R₃ is substituted or unsubstituted (C1-C18) oxime and the R is selected from: acetaldoxime, propionaldoxime, butanaldoxime and benzaldoxime.

94. (Currently amended) A method according to claim 64, wherein at least one R of R₁, R₂ and R₃ is substituted or unsubstituted oxime (C1-C18) and the hydroxylamine is selected from:

Acetaldoxime-3-hydroxylamine,	Butanaldoxime-5-hydroxylamine and
Propionaldoxime-4-hydroxylamine,	(4-benzaldoxime)1-methylhydroxylamine.

95. (Currently amended) A method according to claim 64, wherein at least one R of R₁, R₂ and R₃ is substituted or unsubstituted (C1-C10) hydrazone.

96. (Currently amended) A method according to claim 64, wherein at least one R of R₁, R₂ and R₃ is substituted or unsubstituted (C1-C10) hydrazone and the R is selected from: acetaldehyde hydrazone, propanaldehyde hydrozone, butanaldehyde hydrazone and phenylhydrazone.

97. (Currently amended) A method according to claim 64, wherein at least one R of R₁, R₂ and R₃ is substituted or unsubstituted hydrazone (C1-C10) and the hydroxylamine is selected from

1-hydroxylamine-acetaldehyde hydrazone, 1-hydroxylamine-butanaldehyde hydrazone
1-hydroxylamine-propanaldehyde hydrazone, and
1-hydroxylamine-benzylaldehyde hydrazone.

98. (Currently amended) A method according to claim 64, wherein at least one R of R₁, R₂ and R₃ is substituted or unsubstituted azo.

99. (Currently amended) A method according to claim 64, wherein at least one R of R₁, R₂ and R₃ is substituted or unsubstituted azo and the R is selected from: azobenzene, p-(phenylazo)benzyl and p-diazobenzyl.

100. (Currently amended) A method according to claim 64, wherein at least one R of R₁, R₂ and R₃ is substituted or unsubstituted azo and the hydroxylamine is selected from:

N-(p-phenylazo)benzylhydroxylamine,
N-(p-diazobenzyl)hydroxylamine and
N-(p-methoxylphenylazo)benzylhydroxylamine.

101. (Currently amended) A method according to claim 64, wherein at least one R of R₁, R₂ and R₃ is substituted or unsubstituted (C1-C18) thiol.

102. (Currently amended) A method according to claim 64, wherein at least one R of R₁, R₂ and R₃ is substituted or unsubstituted (C1-C18) thiol and the R is selected from (C1-C18) alkylthiol selected from methyl, ethyl, propyl, butyl, pentyl and hexyl thiol, and arylthiol selected from thiophenol and benzylthiol.

103. (Currently amended) A method according to claim 64, wherein at least one R of R₁, R₂ and R₃ is substituted or unsubstituted (C1-C18) thiol and the hydroxylamine is selected from:

N-(thiomethyl)hydroxylamine,
N-(2-thioethyl)hydroxylamine,

N-(3-thiopropyl)hydroxylamine and
N-(p-sulphydryl)benzylhydroxylamine.

104. (Currently amended) A method according to claim 64, wherein at least one R of R₁, R₂ and R₃ is substituted or unsubstituted (C1-C18) sulfonic acid.

105. (Currently amended) A method according to claim 64, wherein at least one R of R₁, R₂ and R₃ is substituted or unsubstituted (C1-C18) sulfonic acid and the R is selected from methanesulfonic acid, ethanesulfonic acid, propanesulfonic acid, butanesulfonic acid and p-toluenesulfonic acid.

106. (Currently amended) A method according to claim 64, wherein at least one R of R₁, R₂ and R₃ is substituted or unsubstituted (C1-C18) sulfonic acid and the hydroxylamine is selected from:

1-hydroxylamine-methanesulfonic acid, 1-hydroxylamine-butane-4-sulfonic acid and
1-hydroxylamine-ethane-2-sulfonic acid, N-(p-sulfonylbenzyl)hydroxylamine.
1-hydroxylamine-propane-3-sulfonic acid,

107. (Currently amended) A method according to claim 64, wherein at least one R of R₁, R₂ and R₃ is halide.

108. (Currently amended) A method according to claim 64, wherein at least one R of R₁, R₂ and R₃ is halide and the R is selected from F, Cl, Br and I.

109. (Currently amended) A method according to claim 64, wherein at least one R of R₁, R₂ and R₃ is halide and the hydroxylamine is selected from:

N-(chloromethyl)hydroxylamine,
N-(bromomethyl)hydroxylamine,
N-(2-chloroethyl)hydroxylamine,
N-(3-chloropropyl)hydroxylamine,

N-(4-chlorobutyl)hydroxylamine,
N-(p-chlorobenzyl)hydroxylamine,
N-(p-fluorobenzyl)hydroxylamine and
N-(p-iodobenzyl)hydroxylamine.

110. (Currently amended) A method according to claim 64, wherein at least one R of R₁, R₂ and R₃ is substituted or unsubstituted hydroxylamine.

111. (Currently amended) A method according to claim 64, wherein at least one R of R₁, R₂ and R₃ is substituted or unsubstituted hydroxylamine and R is selected from N-methylhydroxylamine, N-ethylhydroxylamine, N-propylhydroxylamine N-butylhydroxylamine, N-pentylhydroxylamine, and N-benzylhydroxylamine.

112. (Currently amended) A method according to claim 64, wherein at least one R of R₁, R₂ and R₃ is substituted or unsubstituted hydroxylamine and the hydroxylamine is selected from:

Bis-methylhydroxylamine,
Bis-(2-ethyl)hydroxylamine,

Bis-(3-propyl)hydroxylamine and
Bis-benzylhydroxylamine.

113. (Currently amended) A method according to claim 64, wherein at least one R of R₁, R₂ and R₃ is substituted or unsubstituted (C1-C18) phosphoester.

114. (Currently amended) A method according to claim 64, wherein at least one R of R₁, R₂ and R₃ is substituted or unsubstituted (C1-C18) phosphoester and the R is selected from: dimethylphosphate, diethylphosphate, dipropylphosphate and benzylphosphate.

115. (Currently amended) A method according to claim 64, wherein at least one R of R₁, R₂ and R₃ is substituted or unsubstituted (C1-C18) phosphoester and the hydroxylamine is selected from:

di-hydroxylaminemethylphosphate ester,
mono-hydroxylaminemethylphosphate ester,

mono-(1-hydroxylamine)-ethyl-2-phosphate ester,
di-(1-hydroxylamine)-2-ethylphosphate ester,
di-(1-hydroxylamine)-3-propyl-phosphate ester,
mono-(hydroxylamine-benzyl-phosphate ester and
di-hydroxylamine-benzylphosphateester.

116. (Previously presented) A method according to claim 62, wherein the method detects the ability of the hydroxylamine to reduce infarct volume in a rat permanent middle cerebral artery occlusion (MCAO) stroke model.

117. (Previously presented) A method according to claim 62, wherein the method detects the ability of the hydroxylamine to protect against amyloid β peptide-induced neuronal cell death.

118. (Previously presented) A method according to claim 62, wherein the method detects the ability of the hydroxylamine to reduce inflammation caused by LPS and INF- γ in E16 rat cortical neuronal cells.

119. (Previously presented) A method according to claim 62, wherein the method detects the ability of the hydroxylamine to reduce β -amyloid-induced increased release of interleukin-1 β in human monocyte THP-1 cells.

120. (Previously presented) A method according to claim 62, wherein the method detects the ability of the hydroxylamine to protect against amyloid β peptide-induced locomotor impairment in rats.

121. (Previously presented) A method according to claim 62, wherein the method detects the ability of the hydroxylamine to reduce spatial learning deficit in rats caused by N-nitro-L-arginine.

122. (Previously presented) A method according to claim 62, wherein the method detects the ability of the hydroxylamine to reduce induction of experimental allergic encephalomyelitis

(EAE) by injection of myelin basic protein (MBP) peptides in rats.

123. (Previously presented) A method according to claim 62, wherein the method detects the ability of the hydroxylamine to reduce weight loss resulting from injection of myelin basic protein (MBP) peptides in rats.

124. (Previously presented) A method according to claim 62, wherein the method detects the ability of the hydroxylamine to reduce acquisition learning deficit in Fas mutated autoimmune mice.

125. (Previously presented) A method according to claim 62, wherein the method detects the ability of the hydroxylamine to reduce neuron loss following brain ischemia and reperfusion injury in gerbils resulting from experimental bilateral carotid occlusion.

126. (Previously presented) A method according to claim 62, wherein the method detects the ability of the hydroxylamine to reduce loss of temporal/spatial short term memory following brain ischemia and reperfusion injury in gerbils resulting from experimental bilateral carotid occlusion.

127. (Previously presented) A method according to claim 62, wherein the method detects the ability of the hydroxylamine to reduce infarct volume following brain ischemia and reperfusion injury in gerbils resulting from experimental bilateral carotid occlusion.

128. (Previously presented) A method according to claim 62, wherein the method detects the ability of the hydroxylamine to reduce lethality volume following brain ischemia and reperfusion injury in gerbils resulting from experimental bilateral carotid occlusion.

129. (Previously presented) A method according to claim 62, wherein the method detects the ability of the hydroxylamine to reduce β AP-mediated inactivation of glutamine synthetase (GS) and creatine kinase (CK) in rat brain tissue extracts and in cultured hippocampal neurons and glia.

130. (Previously presented) A method according to claim 62, wherein the method detects the ability of the hydroxylamine to reduce oxidation-caused side effects of adriamycin anticancer therapy in mice, as measured by acute lethality.

131. (Previously presented) A method according to claim 62, wherein the method detects the ability of the hydroxylamine to delay senescence in IMR90 human lung fibroblasts.